

012103

02570 U.S. PTO

USE OF INHIBITORS OF MAMMALIAN ASPARAGINYL ENDOPEPTIDASE FOR...

Colin Watts

Appl. No.: 09/646,950 Atty Docket: ERP01.004APC

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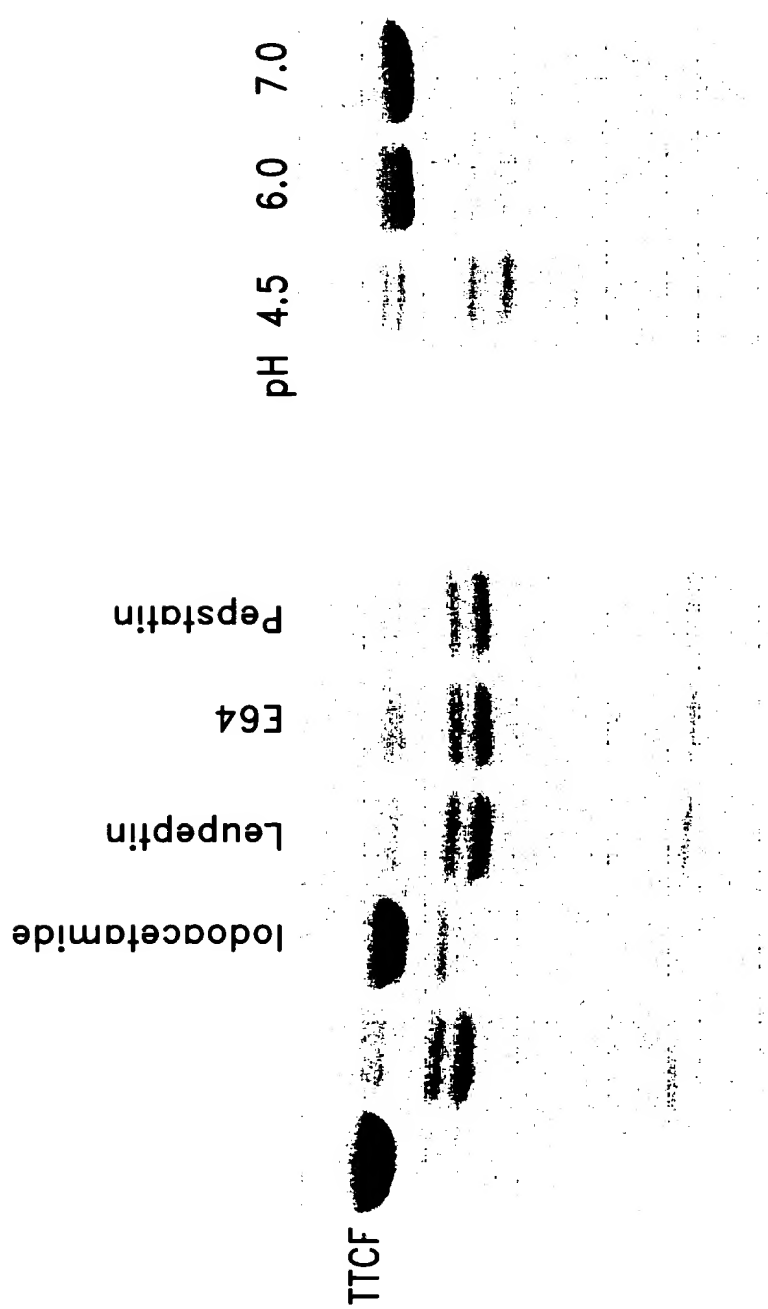


FIG. 1A

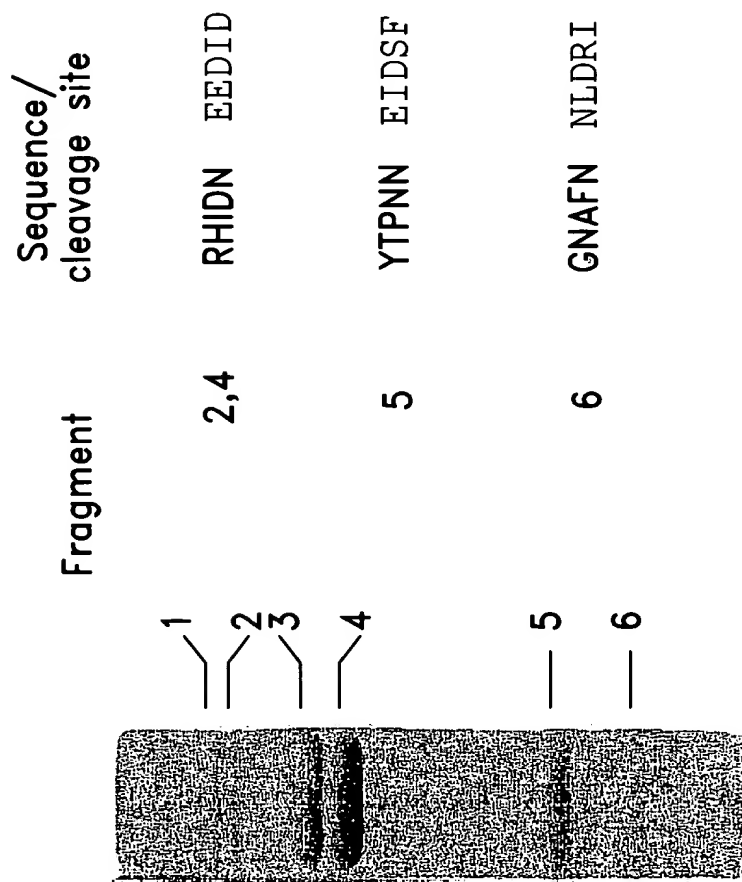


FIG. 1B

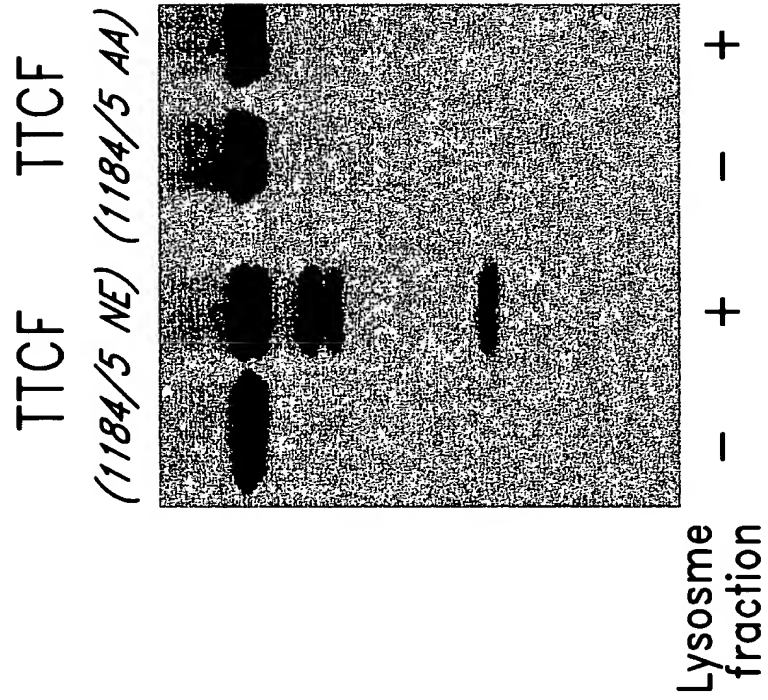


FIG. 1C

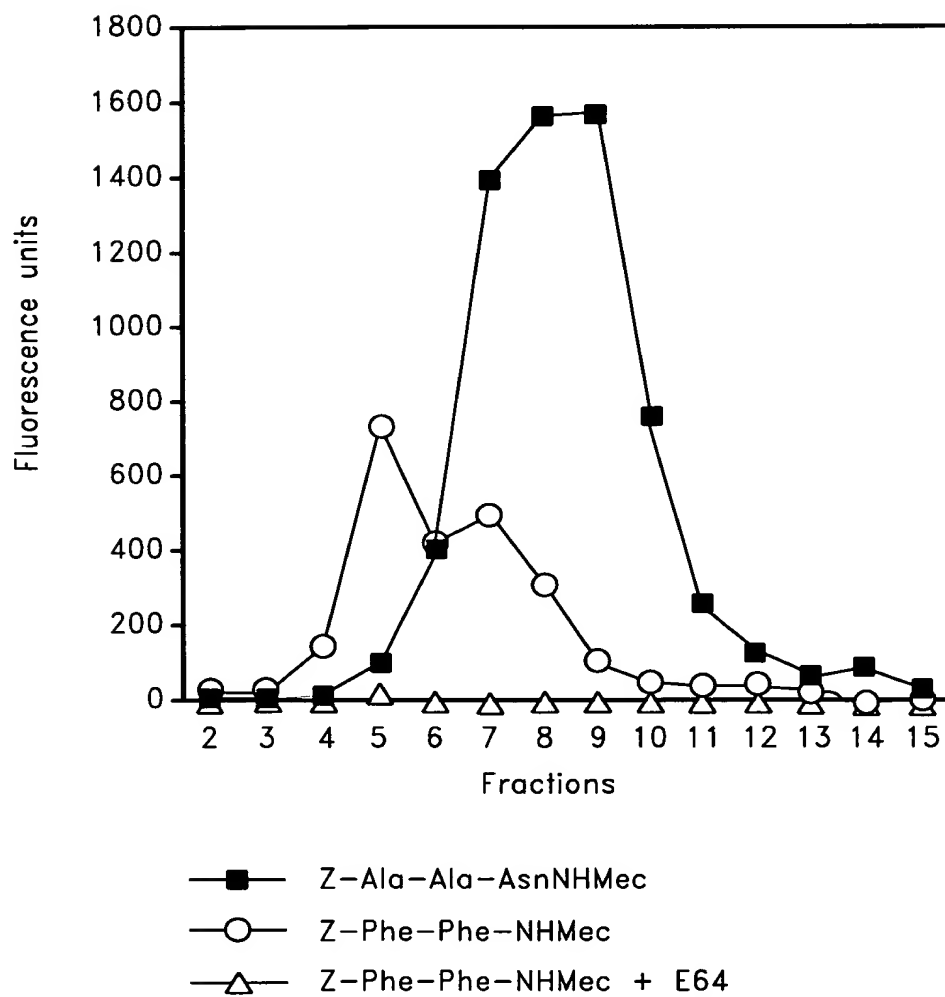


FIG. 2A

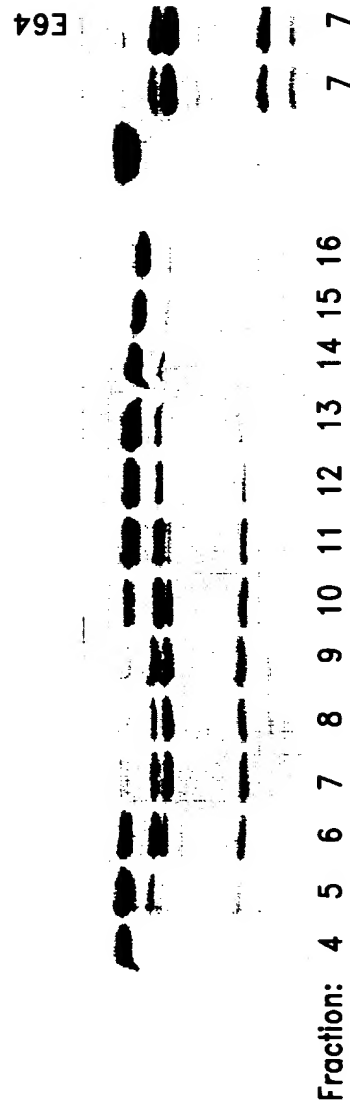


FIG. 2B

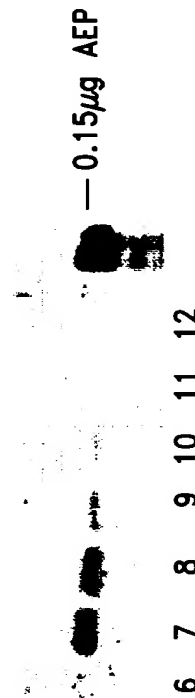


FIG. 2C

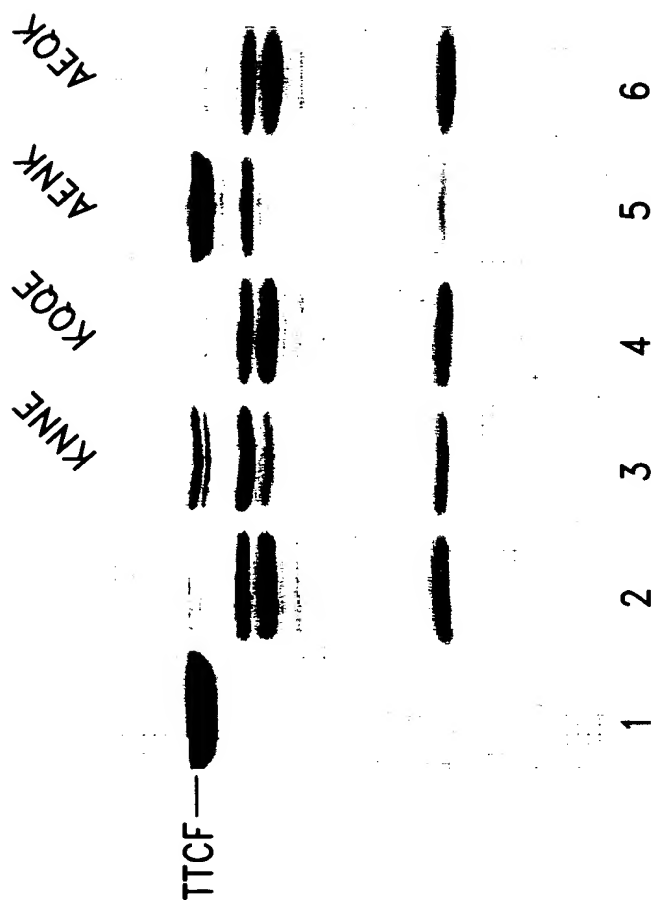


FIG. 3A



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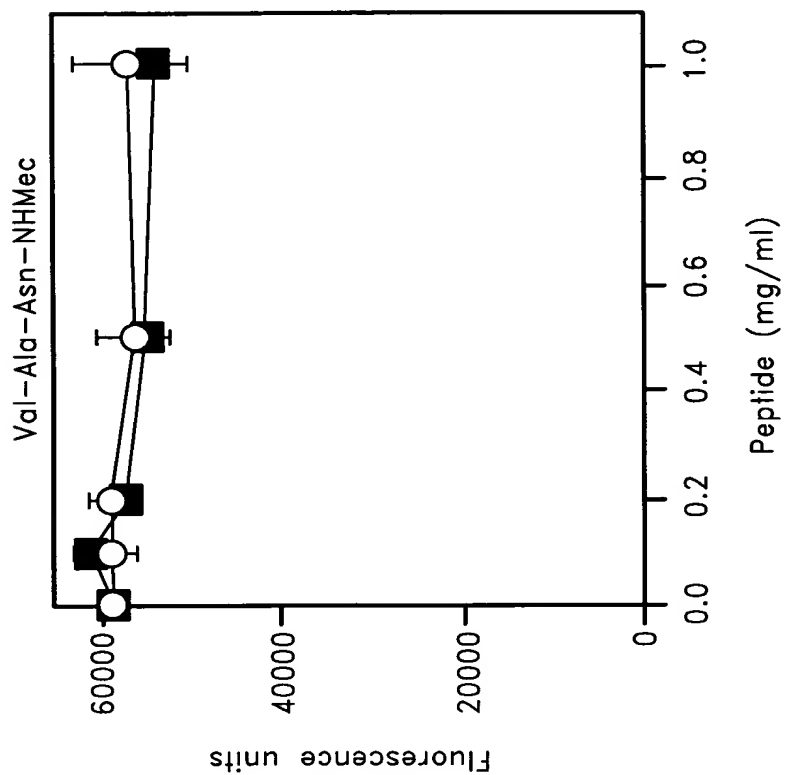
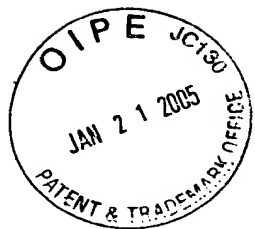


FIG. 3B₂

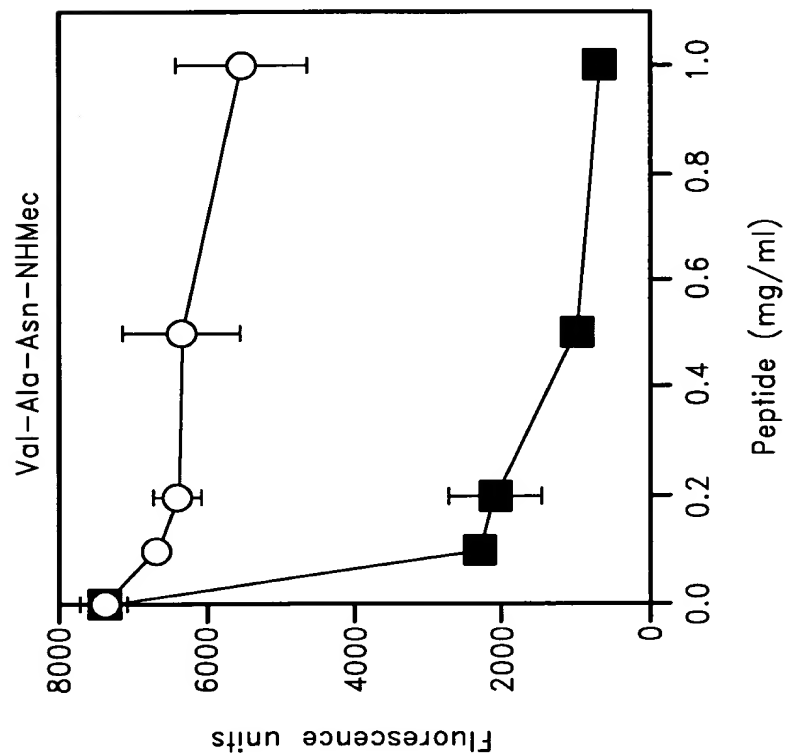


FIG. 3B₁

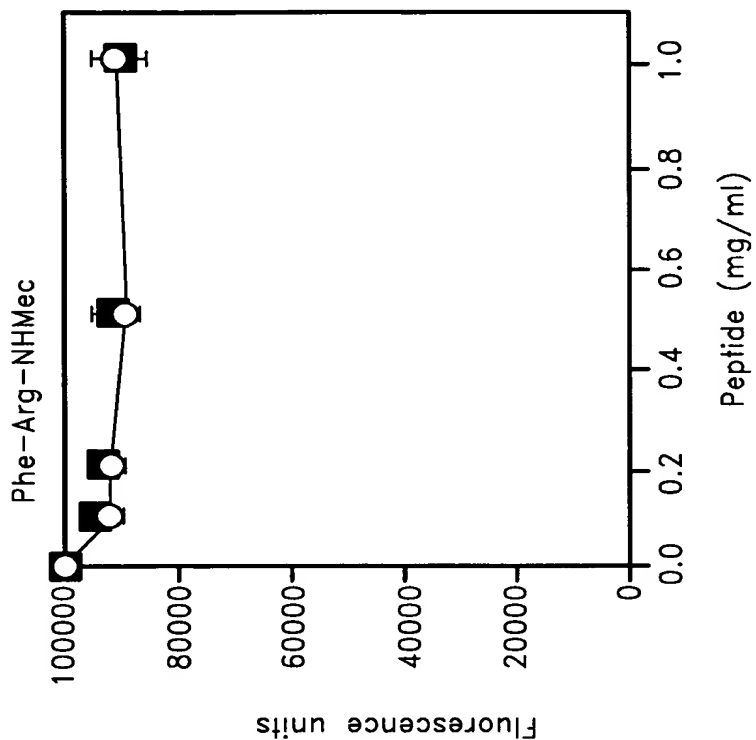
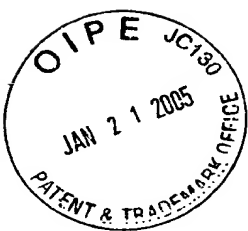


FIG. 3B₄

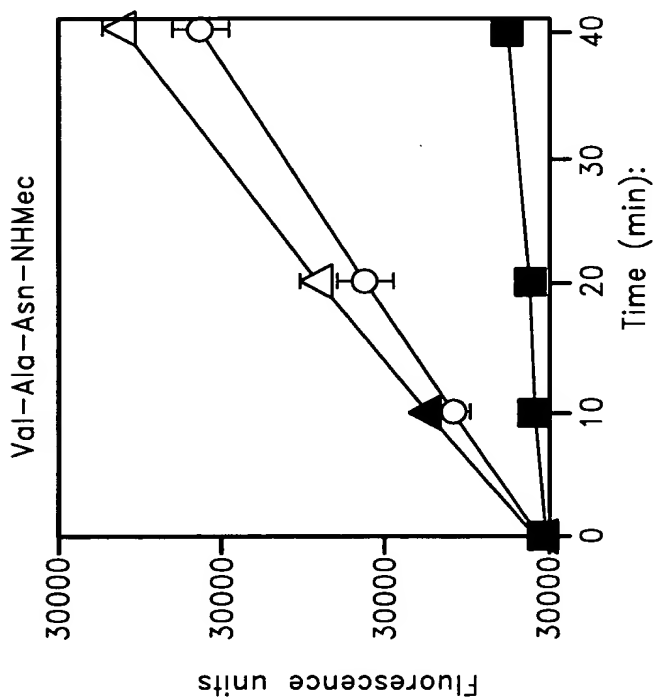
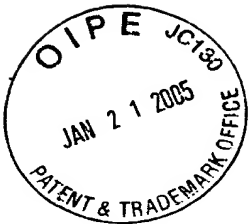
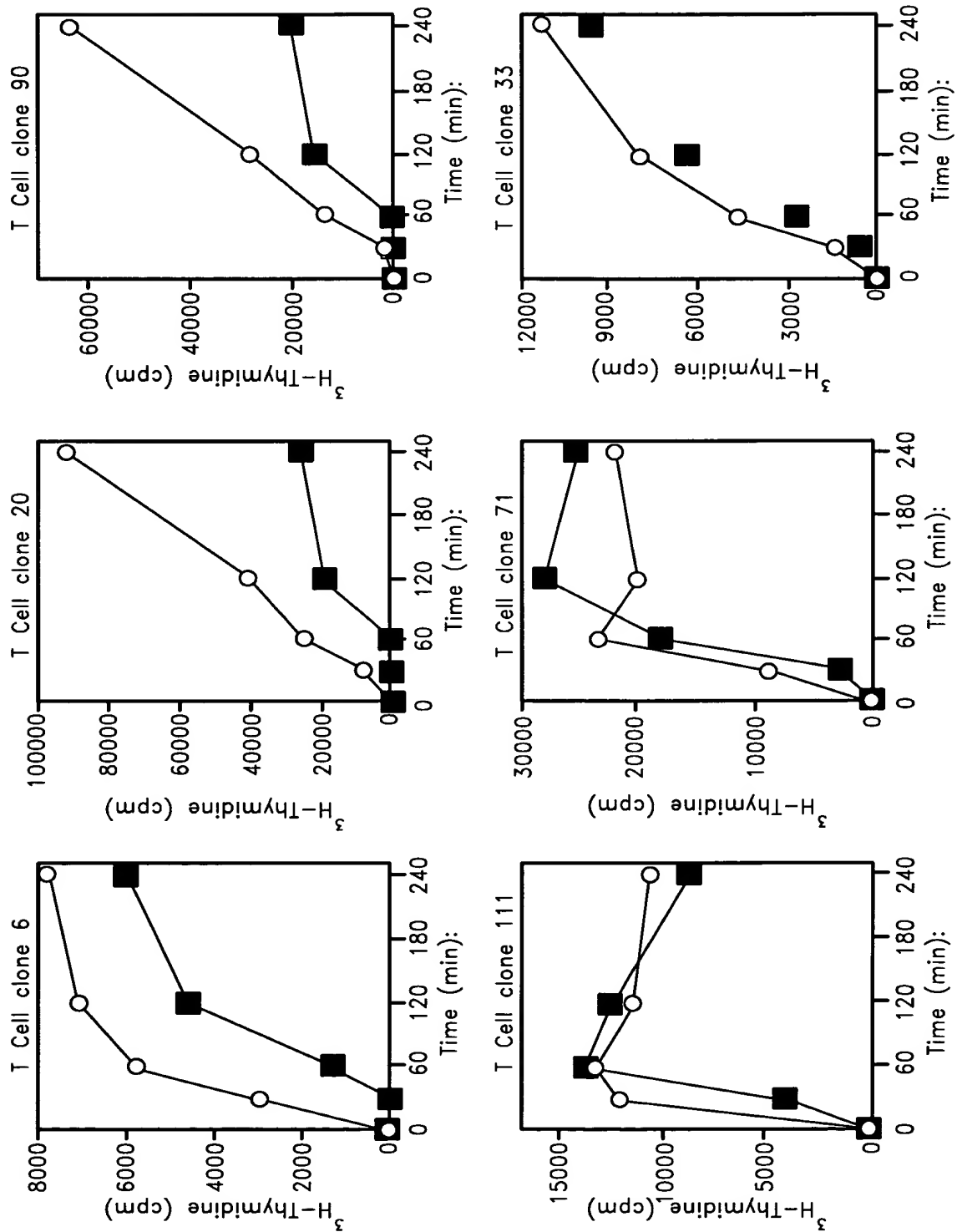


FIG. 3B₃



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FIG. 3C



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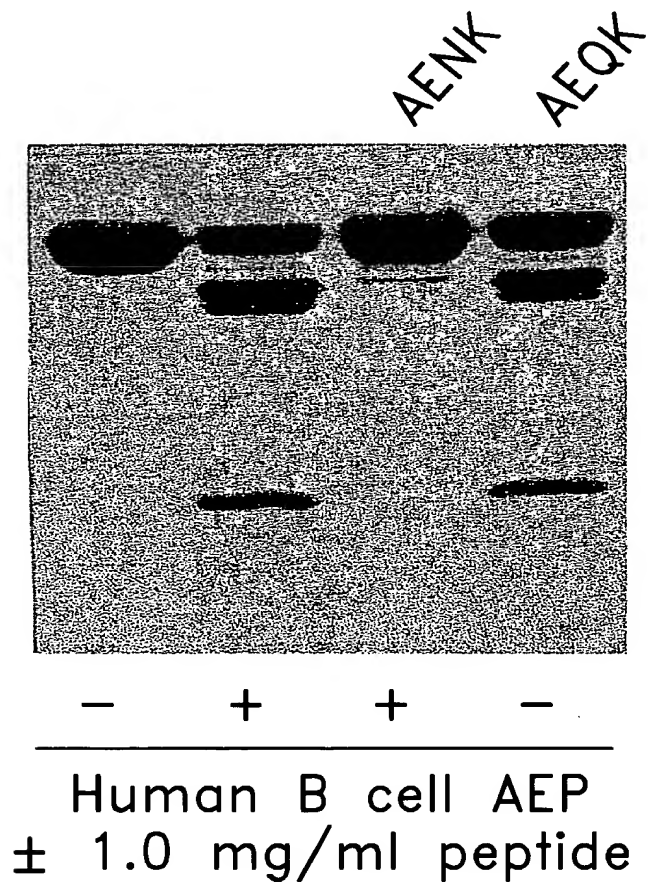


FIG. 3D

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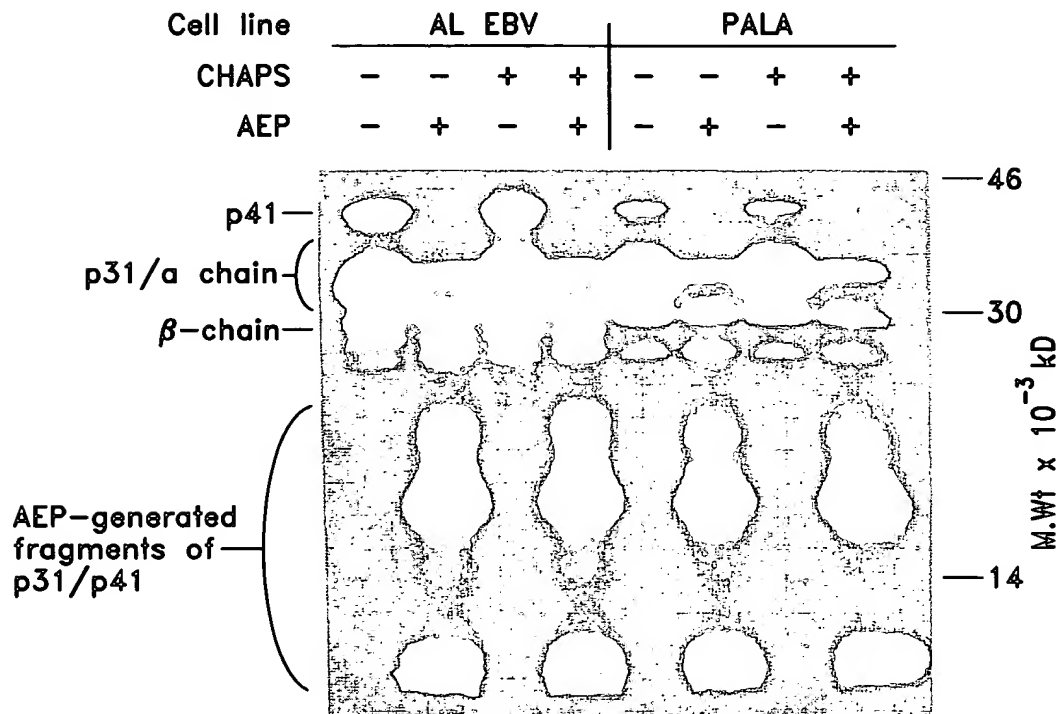


FIG. 4

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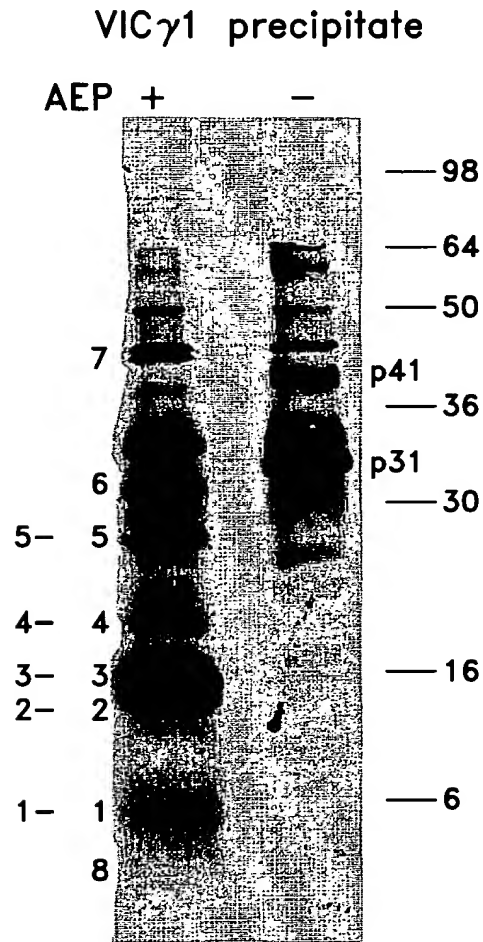
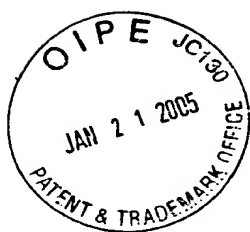


FIG. 5A



N-terminal sequence of bands below

Band 1: N NEQLPM (Peak cycle 5/6)

Band 2: N TMETI (Peak cycle 2)

Band 3: N LRMK (Peak cycle 3)

Band 4: N LRMK
N NEQLPM (Peaks at cycles 3 & 6)

Band 5: N LRMK

Example raw data from band 5

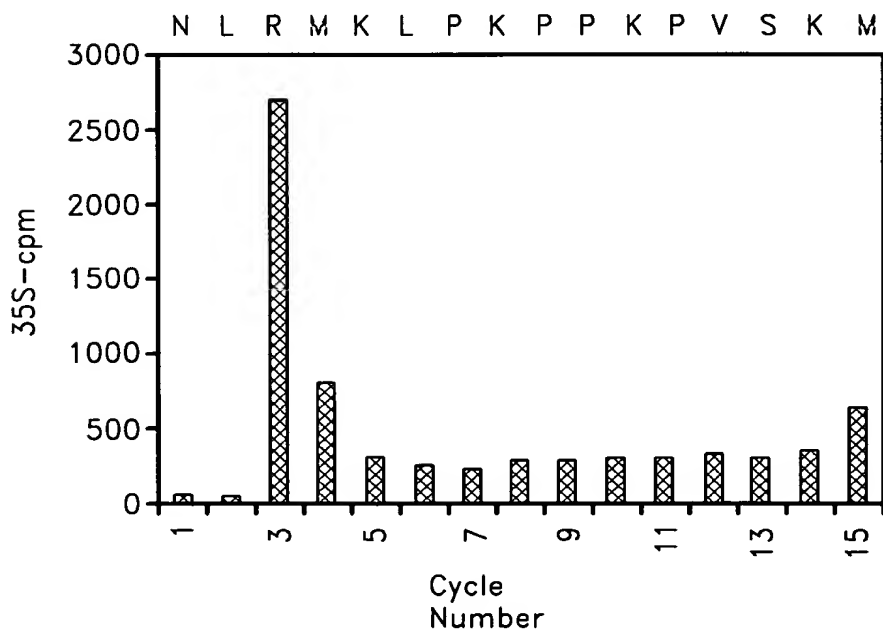


FIG. 5B



↓
MDDQRLDISNNEQLPMLGRRPGAPESKCSRGALYTGSILVTL LLAGQATTAYF
↓
QQQGRLDKLTVTSONLQLENLRMKLPKPPKPVSKMRMATP LLMQALPMGALPQG
↓
QNATKYGNMTEDHVMHLLQNADPLKVYPPLKGSFPENLTHLKN TMETIDWKVFE
MHHWLLFEMSRHSLEQKPTDAPPKESLELEDPS SGLGVTKQDLGPVPM

FIG. 5C



Identified AEP cleavage sites

Tetanus toxin C fragment

RHIDN EEDID
YTPNN EIDSF
GNAFN NLDRI

Ribonuclease

NGQTN CYQSY
VACKN GQTNC

Ovalbumin

GTSVN VHSSL

Hen Egg Lysozyme
(preferred sites listed
first)

GNGMN AWVAW
HGLDN YRGYS
ILQIN SRWWC
VSDGN GMNAW
RWWCN DGRTP
VAWRN RCKGT

Transferrin peptide
(622-642)

LFGSN VTDCS
DCSGN FCLFR

FIG. 6



FIG. 7A

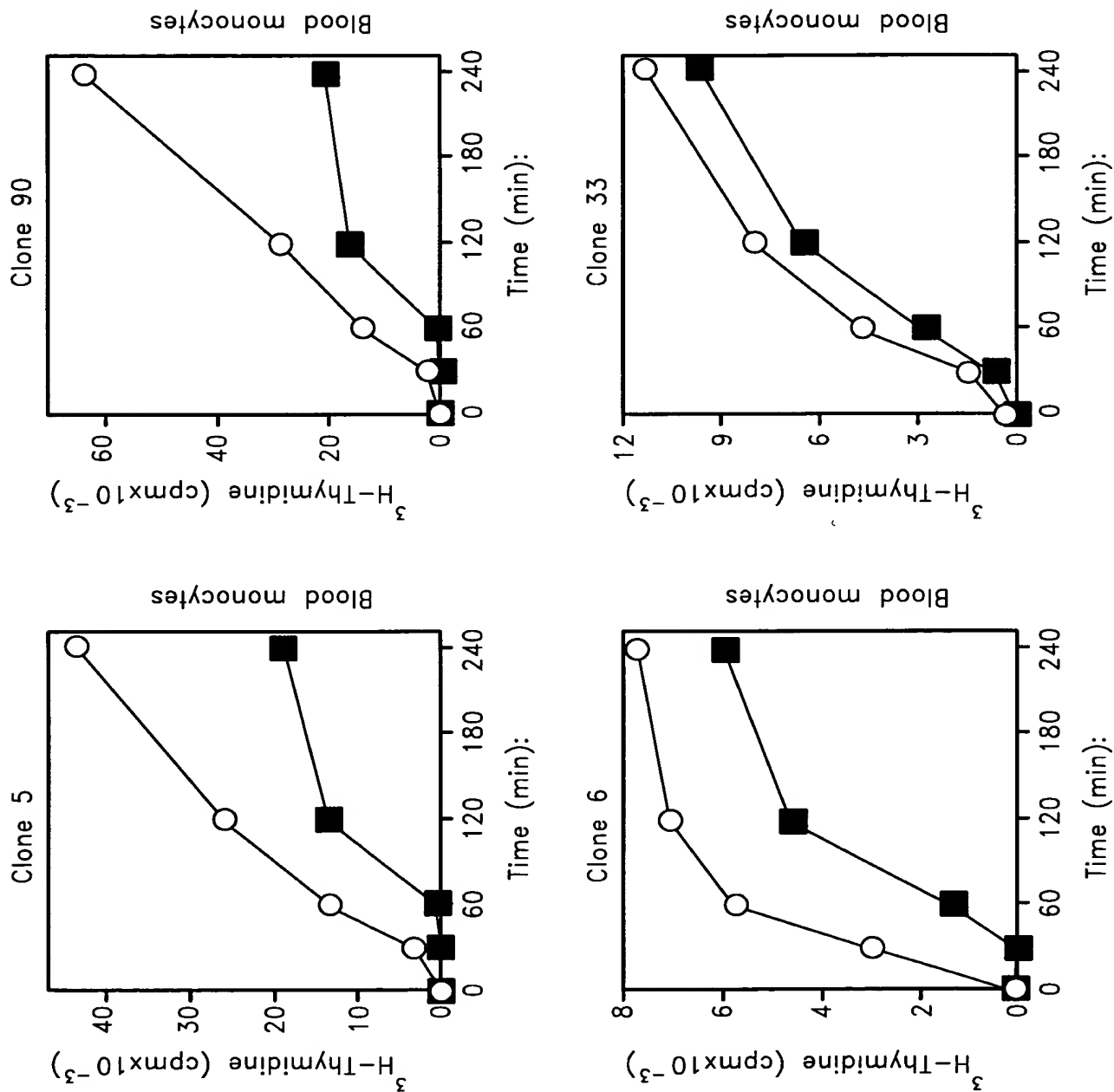
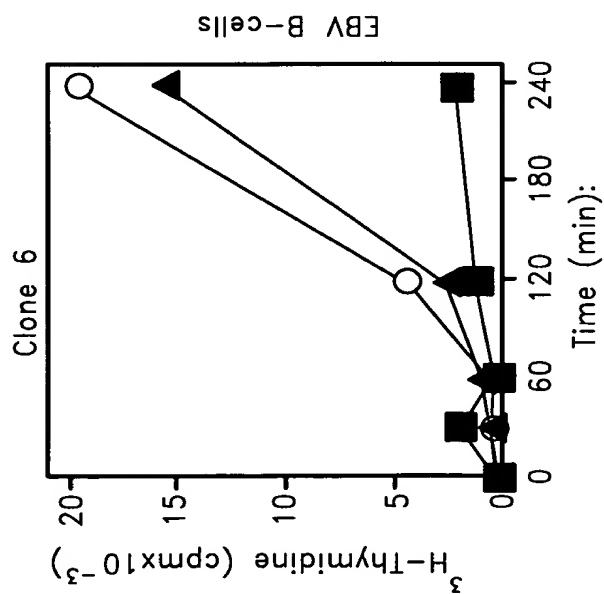
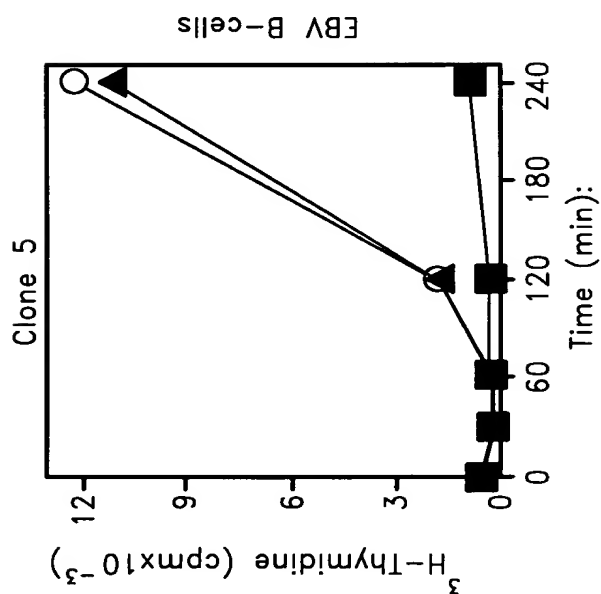
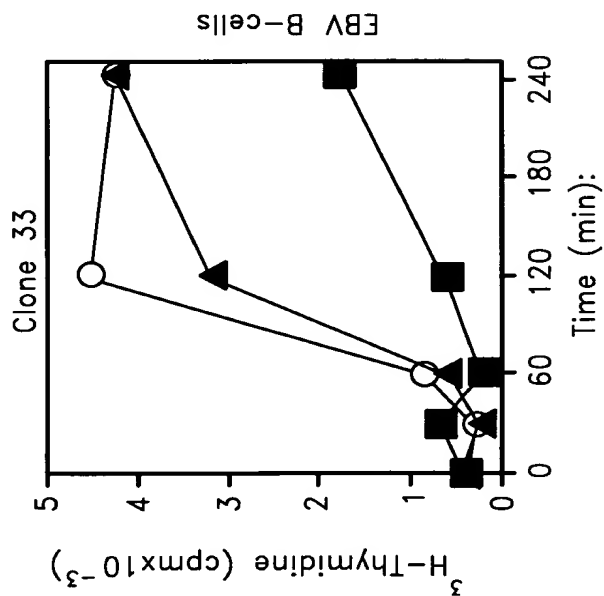
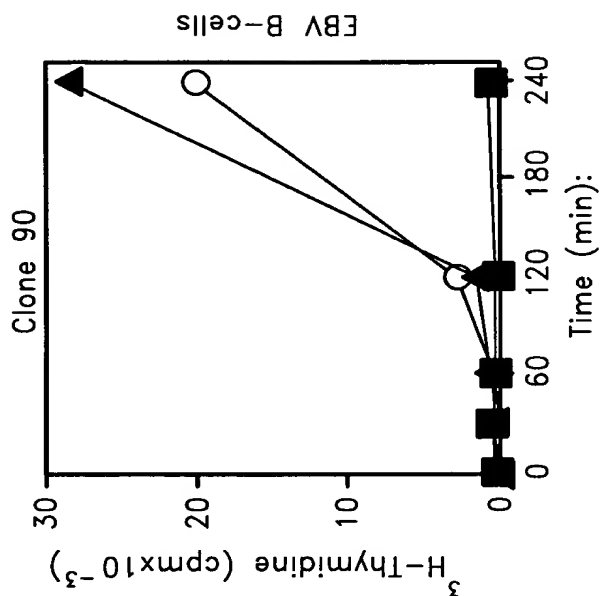




FIG. 7B



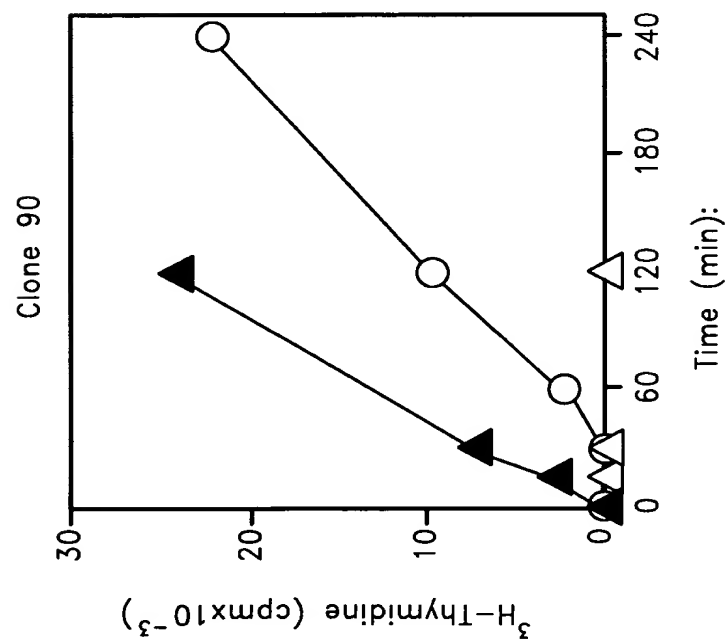
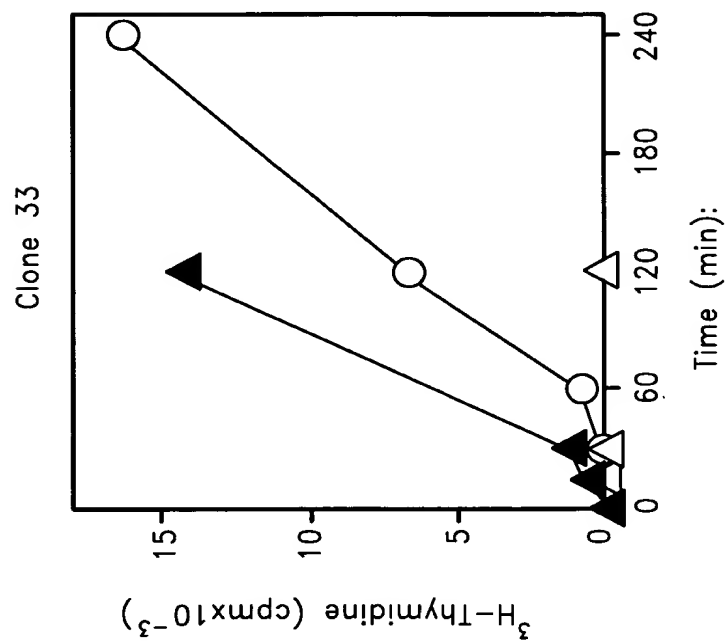


FIG. 7C

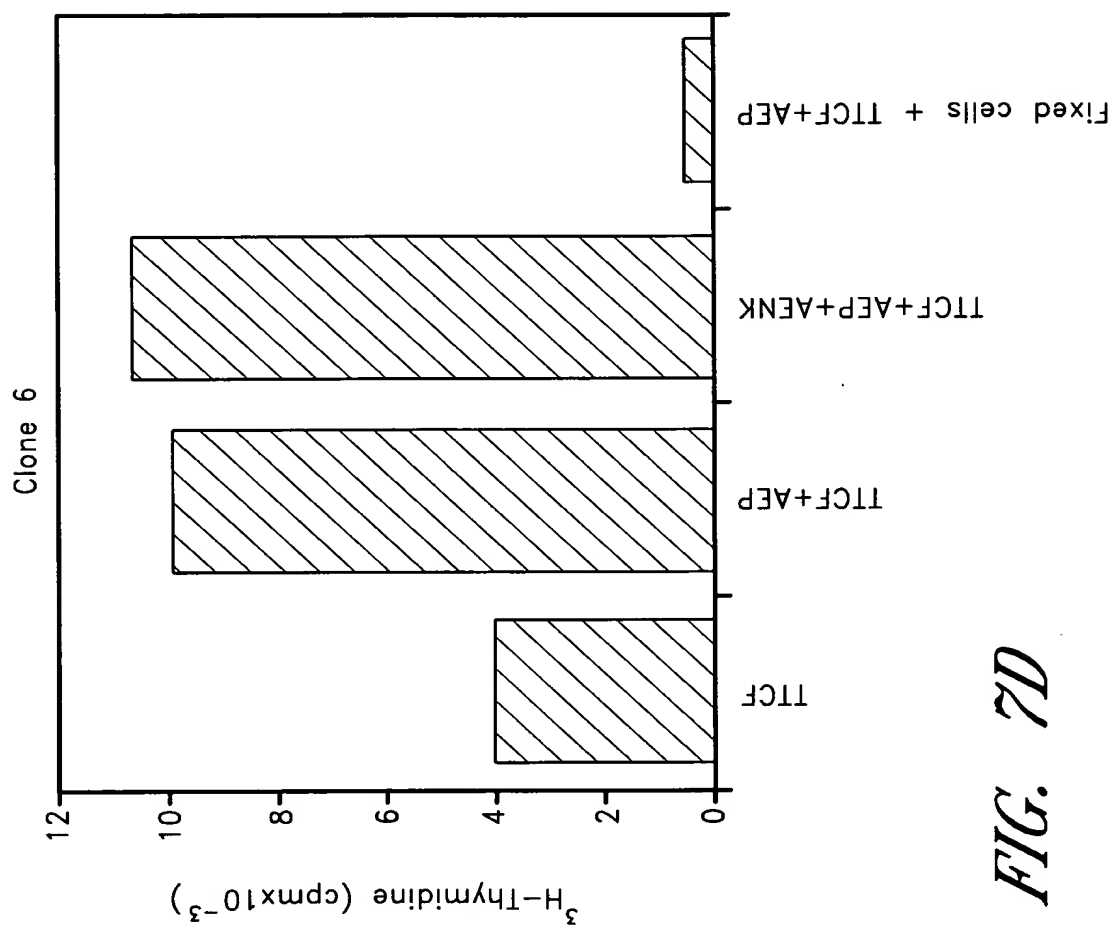


FIG. 7D